Tranexamic Acid



Introduction.

- Tranexamic acid (TXA) is an anti-fibrinolytic and may have immuno-modulatory effects
- TXA been shown to improve survival rates in traumatic haemorrhage and should be administered as soon as possible. Treatment delay reduces the benefit. Every 15 min of treatment delay appears to decrease the relative survival benefit by 10%, with no benefit after 3h.
- The standard dosing regimen is a 1g IV bolus followed by 1g IV over 8 hours. However, there is no evidence of harm with two single bolus doses, and there are likely to be multiple benefits from a 2x 1g bolus regime as the infusion is often omitted or commenced after 3 hours.
- Administering two boluses ensures that 2g will actually be given, removes the need to run an infusion for 8 hours, and allows cognitive offloading of the need for an infusion.
- Increased fibrinolysis is part of traumatic brain injury (TBI) coagulopathy. Haemorrhage expansion occurs in the first few hours after injury and larger hematomas are associated with increased mortality.
- TXA may limit secondary brain injury via inhibition of fibrinolysis (with reduction in ICH progression) and inhibition of tissue plasminogen activator. TXA has been shown to reduce head injury deaths at 24 hours and reduce new haemorrhage formation in those with mild to moderate head injury.

Rationale

2x 1g IV bolus of TXA in traumatic haemorrhage and mild-moderate brain injury will optimize TXA dosing and patient outcomes with no adverse effects.

Safety and clinical governance

All units are encouraged to audit this new regimen to ensure safety and efficacy.

TRAUMATIC CODE RED

Clinical suspicion of severe haemorrhage

AND/OR

At risk of significant haemorrhage*

*Do not delay TXA administration whilst awaiting CT confirmation of cerebral or other haemorrhage or haemodynamic instability. If injury mechanism is commensurate with haemorrhage risk: give 1st dose **TRAUMATIC BRAIN INJURY**

Head Injury + GCS < 12

AND/OR

Head Injury + drop in GCS from 15

AND/OR

Intracranial haemorrhage on CT

AND/OR

HEMS intubated head injury

1g bolus IV pre-hospital AND 1g bolus IV in-hospital

OR

2x 1g bolus IV in hospital

2g IV TOTAL

Initial dose < 1hr from injury

Both doses < 3hr from injury

GCS 13/14 + head injury - prioritise for CT

Aim < 1hr from injury; must be < 3 hours

Adapted from St Georges ED TXA SOP V3.0. Tucker, H, Hudson A, Uprichard J Reviewed M Salmon Sep 2023 Review: Sep 2024

Paediatric Dose (Age < 12 OR weight < 50kg)

- 15mg/kg (max 1g) bolus
- 2mg/kg/hr for 8 hours

References

1. CRASH-2 Trial Collaborators. (2010). Effects of tranexamic acid on death, vascular occlusive events, and blood transfusion in trauma patients with significant haemorrhage (CRASH-2): a randomised, placebocontrolled trial. The Lancet, 376(9734), 23–32. https://doi.org/10.1016/s0140-6736(10)60835-5 2. Morrison, J. J., Dubose, J. J., Rasmussen, T. E., & Midwinter, M. J. (2012). Military Application of Tranexamic Acid in Trauma Emergency Resuscitation (MATTERs) Study. Archives of Surgery, 147(2), 113. https://doi.org/10.1001/archsurg.2011.287

3. July, J., & Pranata, R. (2020). Tranexamic acid is associated with reduced mortality, hemorrhagic expansion, and vascular occlusive events in traumatic brain injury – meta-analysis of randomized controlled trials. BMC Neurology, 20(1), 1–11. https://doi.org/10.1186/s12883-020-01694-4

CRASH-3 Trial Collaborators. (2019). Effects of tranexamic acid on death, disability, vascular occlusive events and other morbidities in patients with acute traumatic brain injury (CRASH-3): a randomised, placebo-controlled trial. The Lancet, 394(10210), 1713–1723. https://doi.org/10.1016/s0140-6736(19)32233-0
Mahmood, A., Needham, K., Shakur-Still, H., Harris, T., Jamaluddin, S. F., Davies, D., Belli, A., Mohamed, F. L., Leech, C., Lotfi, H. M., Moss, P., Lecky, F., Hopkins, P., Wong, D., Boyle, A., Wilson, M., Darwent, M., & Roberts, I. (2020). Effect of tranexamic acid on intracranial haemorrhage and infarction in patients with traumatic brain

injury: a pre-planned substudy in a sample of CRASH-3 trial patients. Emergency Medicine Journal, 0, 1-9. https://doi.org/10.1136/emermed-2020-210424

6. Rowell, S. E., Meier, E. N., McKnight, B., Kannas, D., May, S., Sheehan, K., Bulger, E. M., Idris, A. H., Christenson, J., Morrison, L. J., Frascone, R. J., Bosarge, P. L., Colella, M. R., Johannigman, J., Cotton, B. A., Callum, J., McMullan, J., Dries, D. J., Tibbs, B., ... Schreiber, M. A. (2020). Effect of Out-of-Hospital Tranexamic Acid vs Placebo on 6-Month Functional Neurologic Outcomes in Patients With Moderate or Severe Traumatic Brain Injury. JAMA, 324(10), 961–974. https://doi.org/10.1001/jama.2020.8958

7. Gayet-Ageron A, Prieto-Merino D, Ker K, et al. Effect of Treatment Delay on the Effectiveness and Safety of Antifibrinolytics in Acute Severe Haemorrhage: A Meta-Analysis of Individual Patient-Level Data From 40 138 Bleeding Patients. Lancet. 2018;391(10116):125-132.

8. Guyette, F. X., Brown, J. B., Zenati, M. S., Early-Young, B. J., Adams, P. W., Eastridge, B. J., Nirula, R., Vercruysse, G. A., O'Keeffe, T., Joseph, B., Alarcon, L. H., Callaway, C. W., Zuckerbraun, B. S., Neal, M. D., Forsythe, R. M., Rosengart, M. R., Billiar, T. R., Yealy, D. M., Peitzman, A. B., & Sperry, J. L. (2020). Tranexamic Acid During Prehospital Transport in Patients at Risk for Hemorrhage After Injury. JAMA Surgery, E1–E10. https://doi.org/10.1001/jamasurg.2020.4350

9. Drew, B., Auten, J., Cap, A., Deaton, T., Donham, B., Dorlac, W., DuBose, J., Fisher, A. D., Ginn, A. J., Hancock, J., Holcomb, J. B., Knight, J., Knight, R., Koerner, A., Littlejohn, L., Martin, M. J., Morey, J., Morrison, J., Schreiber, M., ... Butler, F. (2020). The Use of Tranexamic Acid in tactical combat casualty care. Journal of Special Operations Medicine, 20(3), 36–43. https://doi.org/10.0000/0000